In The Claims

Amend Claims 2 and 65, and add Claim 106 as shown in the enclosed listing.

1. (cancelled)

- 1 2. (currently amended): A method of treating degenerative diseases due to
- 2 acquired mitochondrial DNA damage;
- 3 redox damage to mitochondrial macromolecules
- and inherited mitochondrial genetic defects
- said method comprising the steps of: selecting a <u>non-superoxide dismutase mimic</u> composition
- from a group consisting of open ring polyamines, macrocyclic polyamines, branched linear
- 7 polyamines and substituted polyamines;
- 8 synthesizing said composition; and
- administering an effective dose of said composition to a mammal;
- wherein said step of synthesizing comprises converting by treatment with an alkyl halide a
- compound taken from a group consisting of those compounds having the formula

- wherein A and B are hydrogen or alkyl, and m,n, and p are the same or different, and those
- compounds having the formula

NH HN

3. (original): The method of claim 2 wherein said composition is taken from a group consisting of those compositions having the formulae:

4 and

$$R_1$$
 R_2 R_3

6 wherein:

 R_1 and R_2 are taken from a group consisting of hydrogen, alkyl, aryl, cycloalkyl, amino acid, glutathione, uric acid, ascorbic acid, taurine, estrogen, dehydroepiandrosterone, probucol, vitamin E, hydroxytoluene, carvidilol, α -lipoic acid, α -tocopherol, ubiquinone, phylloquinone, β -carotene, meanadione, glutamate, succinate, acetyl-L-carnitine, co-enzyme

- Q, lazeroids, polyphenolic flavonoids, homocysteine, menaquinone, idebenone, dantrolene,
- $-(CH_2)_n[XCH_2)_n]NH_2$ wherein n = 3-6 and R_1 and R_2 taken together are $-(CH_2XCH_2)_n$
- 13 wherein n = 3-6,
- R₃ and R₄ are taken from a group consisting of hydrogen, alkyl, aryl, cycloalkyl, amino acid,
- glutathione, uric acid, ascorbic acid, taurine, estrogen, dehydroepiandrosterone, probucol,
- vitamin E, hydroxytoluene, carvidilol, α-lipoic acid, α-tocopherol, ubiquinone,
- 17 phylloquinone, β-carotene, meanadione, glutamate, succinate, acetyl-L-carnitine, co-enzyme
- Q, lazeroids, polyphenolic flavonoids, homocysteine, menaquinone, idebenone, dantrolene or
- heterocycle and R_3 and R_4 taken together are $-(CH_2XCH_2)_n$ wherein n = 3-6,
- 20 R₅ and R₆ are taken from a group consisting of hydrogen, alkyl, aryl, cycloalkyl, amino acid,
- glutathione, uric acid, ascorbic acid, taurine, estrogen, dehydroepiandrosterone, probucol,
- vitamin E, hydroxytoluene, carvidilol, α -lipoic acid, α -tocopherol, ubiquinone,
- phylloquinone, β-carotene, meanadione, glutamate, succinate, acetyl-L-carnitine, co-enzyme
- Q, lazeroids, polyphenolic flavonoids, homocysteine, menaquinone, idebenone, dantrolene
- $-(CH_2)_n[XCH_2)_n]NH_2$ wherein n = 3-6, and R₅ and R₆ taken together are $-(CH_2XCH_2)_n$
- 26 wherein n = 3-6.
- 27 M, n, and p may be the same or different and are bridging groups of variable length from 3-12
- carbons, and
- 29 X is taken from a group consisting of nitrogen, sulfur, phosporous and carbon.
- 4. (Withdrawn): The method of Claim one wherein said step of synthesizing further comprises
- the steps of:
- 3 -admixing an element taken from a group consisting of 2,4 dibromopropane and absolute
- 4 ethanol into 1,2-diaminoethane hydrate;
- 5 -heating the resulting mixture to approximately 50°C for about one hour;

- 6 -adding potassium chloride;
- 7 -continuing said heating for three hours;
- 8 -filtering potassium bromide out of the mixture;
- 9 -distilling the mixture at reduced pressure;
- -allowing the formation of top and bottom layers;
- -separating and distilling the top layer;
- -converting free amine in the distilled top layer to a tetrahydrochloride salt; and
- -converting said salt to a free amine by treatment with ammonium hydroxide.
- 1 5. (Withdrawn): The method of claim 4 wherein said step of converting to a
- 2 tetrahydrochloride salt comprises adding hydrochloric acid to said distilled top layer.
- 6. (original): The method of Claim 4 wherein said composition consists of 1,3-bis-[(2'-
- aminoethyl)-amino]propane and step of admixing a solution comprises preparing said solution
- by mixing 1,3-dibromopropane and absolute ethanol in a ratio of approximately 1 to 3 per
- 4 weight.
- 1 7. (Withdrawn): The method of Claim 6 wherein said step of admixing further comprises
- 2 slowly adding said solution into 1,2-diaminoethane hydrate in a ratio of approximately 2.6 to 1
- 3 per weight.
- 1 8. (Withdrawn): The method of claim 7 wherein, the step of preparing said solution
- 2 comprises mixing 15 grams of 1,3-diaminopropane and 50 milliliters of absolute ethanol; and
- 3 the step of slowly adding comprises adding said solution to 20 grams of potassium chloride;

- 9. (Withdrawn): The method of Claim 8 wherein said step of converting to a tetrahydrochloride salt comprises adding six molar concentration of hydrochloric acid.
- 10.-13. (canceled)

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- 1 14. (Withdrawn): The method of Claim 13 wherein said degenerative diseases comprise neurodegenerative diseases characterized by excess iron pools and said compound is selected from a group consisting of 2,2,2-piperidine and 2,3,2 adamantane.
- 1 16. (Withdrawn): The method of Claim 13 wherein said degenerative diseases comprise
 2 neurodegenerative diseases and strokes; and said composition is selected from a group
 3 consisting of compositions having open ring metal binding molecules taken from a group
 4 consisting of compositions having copper binding molecules and manganese binding
 5 molecules.
- 1 17. (Withdrawn): The method of Claim 16 wherein said compositions having copper-binding molecules include 2,3,2 isopropyl on N1/N4; and

said compositions having manganese-binding molecules include 3,3,3 tetramine.

18.-24. (canceled)

- 25. (Withdrawn): The method of Claim 22 wherein said degenerative disease comprises
 Alzheimer's disease and presbycussis; and
 said composition is derived from compounds selected from a group consisting of α lipoic acid
- 4 and acetyl-l-carnitine polyamines.

26-28. (canceled).

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1 29. (Withdrawn): The method of Claim 22 wherein said degenerative diseases

comprise cancer; and said composition is taken from a group consisting of cobalt di-

homocysteine polyamines.

30.-37. (cancelled)

38. (Withdrawn): The method of Claim 20 wherein;

said compound consisting of pyridine tetramine.

39.-43. (canceled)

- 1 44. (Withdrawn): The method of Claim 4 wherein said composition consists of (2-
- aminoethyl){3-[(2-aminoethyl)amino]-1-methylbutyl}amine; and said step of admixing a
- 3 solution comprises preparing said solution by mixing 2,4 dibromopropane and absolute
- 4 ethanol in a ratio of approximately 1 to 20 per weight.
- 1 45. (Withdrawn): The method of claim 44 wherein said step of admixing comprises slowly
- adding said solution into 1,2-diaminoethane hydrate in a ratio of approximately 44 to 1 per
- weight.
- 1 46. (Withdrawn): The method of claim 45 wherein said step of converting to a
- tetrahydrochloride salt comprises of adding hydrochloric acid.

- 1 47. (Withdrawn): The method of Claim 2 wherein said composition consists of (2
 - aminoethyl){3-[(2-aminoethyl)amino]-1-methylbutyl}amine; and
 - said step of synthesizing further comprises; the steps of
 - -admixing a solution of an element, taken from a group consisting of 1,3-diaminopropane and
 - N,N-dimethyl-1,3-propanediamine and ethanol into 2-chloromethylpiperidine in water;
 - -adjusting the pH of the resulting mixture to 9 by addition of 10% sodium hydroxide;
 - -stirring the mixture at room temperature and maintaining the pH between 8 and 9 by addition
 - 8 of sodium hydroxide over 3 days;
 - 9 -allowing solvents to evaporate; and
 - -extracting residues with CH₂Cl₂.
 - 1 48. (Withdrawn): The method of Claim 47 wherein said step of admixing a solution further
 - 2 comprises adding said solution into chloromethyl pyridine in water in a ratio of approximately
 - 5 to 3 per weight wherein said chloromethylpyridine is diluted into water in a ratio of
 - approximately 1 to 5 per weight.
 - 1 49. (Withdrawn): The method of claim 48 wherein said step of admixing a solution
 - comprises preparing said solution in a ratio of approximately 1 to 50 per weight.
 - 1 50. (Withdrawn): The method of Claim 49 wherein said steps of synthesizing comprises
 - 2 synthesizing
 - 3 (2-pyridylmethyl){3-[(2-pyridylmethyl)amino]propyl}amine; and
 - said step of admixing a solution further comprises preparing said solution by mixing 1,3-
 - 5 diaminopropane in water with ethanol.

- 51. (Withdrawn): The method of claim 50 when said step of synthesizing further comprises 1 synthesizing methyl(3-[methyl(2-pyridylmethyl)amino]propyl}(2-pyridylmethyl)amine; and 2 said step of admixing a solution further comprises preparing said solution by mixing N,N-3 dimethyl-1,3 propanediamine in water with ethanol.
- The method of claim 2 wherein said step of synthesizing comprises the 52. (Withdrawn): 1 steps of a preparation by adding a first solution of 1,3 diaminopropane and absolute ethanol 2 dropwise into a second solution of ethanol and an element taken from a group consisting of 1-3 (2chloroethyl)piperidine and 1-(2-chloroethylpiperizine) and admixing over approximately 30 4 minutes; 5
- 6 stirring said preparation over approximately 24 hours;
- evaporating the solvents in said preparation; 7
- extracting the residue using a volume of CH₂Cl₂ dried over Na₂SO₄ and evaporated to dryness; 8
- purifying the resulting composition by converting to its hydrochloride salt by adding 9
- hydrochloric acid; and 10

- converting said salt to its free amine by treatment with NH₄OH. 11
- 53. (Withdrawn): The method of claim 52 wherein said step of mixing a preparation 1 comprises 2
- forming said first solution of 1,3 diaminopropane and ethanol in a ratio of approximately 1 to 3
- 100 per weight and adding said first solution into said second solution in a ratio of 4
- approximately 1 to 1 by weight. 5

- 1 54. (Withdrawn): The method of Claim 2 wherein said composition consists of
- 2 [2-(methylethylamino)ethyl](3-{[2-(methylamino)ethyl]amino}propyl)amine; and said step of
- 3 synthesizing further comprises; preparing of first mixture of magnesium turnings,
- 4 1,3-bis-[(2'-aminoethyl)-amino]propane, benzene and acetyl chloride respective approximate
- 4 percentage of 0.6%, 8.5%, 8.45%, and 6.4% per weight;
- 5 cooling said first mixture;
- separating the mixture into a liquid phase and a solid phase;
- 7 preparing a second mixture by mixing said solid phase with ether;
- preparing a solution by pouring said second mixture over ice;
- preparing a third mixture by adding said solution to said liquid phase;
- washing said third mixture with sodium bicarbonate;
- washing said third mixture with water.
- 1 55. (Withdrawn): The method of Claim 2 wherein said step of synthesizing comprises
- 2 converting the starting di or tetramine component, at least one of said components in said
- 3 compounds to the corresponding N-substituted compound by treatment with an alkyl halide;
- 4 and
- 5 purifying said composition by conversion to a salt through addition of hydrochloric acid.
- 1 56. (Withdrawn): The method of Claim 2 wherein said composition consists of (2-
- aminoethyl){3-[(2-aminoethyl)methylamino]propyl}methylamine, and
- 3 said step of synthesizing further comprises:
- preparing a first solution of N,N-dimethyl-1,3-propanediamine and ethanol in a ratio of
- 5 approximately 1 to 50 per weight;
- preparing a second solution of 2-chloroethylamine and ethanol in a ratio of approximately 1 to
- 7 17 per weight;

- 8 combining said first and second solutions into a third solution;
- 9 stirring said third solution at room temperature for approximately 20 hours;
- evaporating solvents in said third solution; and
- extracting residues in said solution with a volume of CH₂Cl₂.
- 1 57. (Withdrawn): The method of Claim 2 wherein said composition consists of
- 2 [2-(bicyclo[3.3.1]non-3-ylamino)ethyl](3-{2-(bicyclo[3.3.1]non-3-
- 3 ylamino)ethyl]amino}propyl)amine, and said step of synthesizing further comprises heating
- for approximately 6 hours at 215°C a mixture of 1-bromoadamantane and 2,3,2-tetramine in a
- 5 mol ratio of approximately 1 to 5;
- admixing said mixture into a solution of 2NHCl and ether having a ratio of approximately 1.25
- to 1 per weight, in a ratio of approximately 1 to 9 per weight;
- separating the aqueous layer and alkalinizing said layer in a volume of 50% aqueous NaOH;
- 9 extracting with ether;
- drying the extract over K_2CO_3 ; and
- evaporating to an oil.
- 1 58. (Withdrawn): The method of Claim 2 wherein said composition consists of [2-
- 2 (methylethylamino)ethyl](3 {[2-(methylamino)ethyl]amino}propyl)amine; and
- 3 said methylating step of synthesizing further comprises;
- 4 methylating terminal nitrogens of 2,3,2 tetramine by refluxing in the presence of benzene and
- 5 acetyl chloride.
- 1 59. (Withdrawn): The method of Claim 58 wherein said step of synthesizing further
- 2 comprises;
- 3 preparing a first mixture of magnesium turnings;

- of 1,3-bis-[(2'-aminoethyl)-amino]propane, benzene and acetyl chloride respective
- approximate percentage of 0.6%, 8.5%, 8.45%, and 6.4% per weight;
- 6 cooling said first mixture;
- separating the mixture into a liquid phase and a solid phase;
- 8 preparing a second mixture by mixing said solid phase with ether;
- 9 preparing a solution by pouring said second mixture over ice;
- preparing a third mixture by adding said solution to said liquid phase;
- washing said third mixture with sodium bicarbonate;
- washing said third mixture with water;
- drying said third mixture over CaCl₂;
- filtering said third mixture;
- preparing a fourth mixture of said third mixture sodium hydride and N,N,-dimethylformamide
- in a ratio of approximately 2.5, 1 and 37.5 respectively per weight;
- heating said fourth mixture under N₂ at approximately 60°C for about three hours;
- treating said fourth mixture with approximately ¼ its volume of iodomethane;
- stirring said treated fourth mixture at 50°C for approximately 24 hours;
- quenching said treated fourth mixture with 95% ethanol;
- removing volatiles at reduced pressure;
- watering with addition of approximately ½ volume of water;
- extracting organic products with approximately three ½ volumes of chloroform;
- washing said organic products with water and NaCl;
- drying said organic products over anhydrous sodium sulfate;
- concentrating into an oil;
- purifying said oil by flash chromatography with ¼ hexanes-ethyl acetate as eluent into an
- acetylated oil of said composition;

- forming a solution of said acetylated oil, potassium hydroxide, methanol and water in
- respective proportions of 1, 3, 23 and 5 per weight respectively;
- 31 heating said solution under reflux for about 24 hours;
- removing methanol at reduced pressure;
- extracting into ether;
- washing with NaCl;
- 35 drying over sodium sulfate;
- 36 concentrating under vacuum;
- purifying by flash chromatography; and
- 38 evaporating solvents.
- 1 60. (Withdrawn): The method of Claim 2 wherein said composition consists of [2-
- 2 (dimethylamino)ethyl](3-{[2-(dimethylamino)ethyl]methylamino}propyl)methylamine; and
- 3 said steps of synthesizing further comprises;
- refluxing for about 20 hours a solution of 2,3,2 tetramine, formic acid and 37% formaldehyde
- and water in a weight proportions of approximately 1,10,10 and 1 respectively;
- 6 evaporating solvents from said solution;
- 7 making said solution basic by addition of NaOH; and
- 8 extracting residues with 3 times $1\frac{1}{2}$ volume of CH_2Cl_2 .
- 1 61. (Withdrawn): The method of Claim 2 wherein said composition consists of 2-[3-(2-
- aminoethylthio)propylthio]ethylamine; and
- said step of synthesizing further comprises:
- preparing a first solution of 1,3-dimercaptopropane and water in a weight ration of about 1 to
- 5 50;
- 6 preparing a second solution of NaOH and water in a weight ratio of about 1.5 to 10;

- g 1;
- forming a third solution of 2-chloroethylamine and ethanol in a weight ratio of about 8.5 to 1;
- admixing said solution into said mixture in a ratio of about 1 to 3.8;
- refluxing said mixture over approximately 8 hours;
- evaporating solvents from said refluxed mixture;
- extracting residues with CH₂Cl₂.
 - 62-64. (cancelled)
- 65. (currently amended) A method of treating degenerative diseases due to
- 2 <u>acquired mitochondrial DNA damage</u>;
- 3 redox damage to mitochondrial macromolecules
- 4 and inherited mitochondrial genetic defects
- 5 said method comprising the steps of: selecting a non-superoxide dismutase mimics
- 6 composition from a group consisting of open ring polyamines, macrocyclic polyamines,
- 7 <u>branched linear polyamines and substituted polyamines;</u>
- 8 synthesizing said composition; and
- administering an effective dose of said composition to a mammal;
- wherein said step of synthesizing comprises converting by treatment with an alkyl halide a
- compound taken from a group consisting of those compounds having the formula:

wherein A and B are hydrogen or alkyl, and m,n, and p are the same or different, and those compounds having the formula:

NH HN

- 16 [The method of Claim 2] wherein said composition consists of 1,4,8,11-tetraaza-1,4,8,11-
- tetraethylcyclotetradecane; and

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- said step of synthesizing further comprises:
- forming a solution of cyclam and DMF in a weight ration of approximately 1- to 50;
- admixing under stirring small portions of NaH in a weight ratio of about 1 to 12.5;
- 21 Heating said solution for about three hours at about to degrees C;
- admixing iodoethane in a single portion into said solution in a weight ratio of about 1 to 17.5;
- Heating said solution at about 60 degrees C over about 18 hours;
- quenching the solution with about 95% ethanol;
- extracting residue with CH₂CH₂.
- 1 66. (Withdrawn): The method of Claim 2 wherein said composition consists of N,N'-(2'
- dimethylphosphinoethyl)-propylenediamine; and the step of synthesizing further comprises:
- 3 incorporating phosphorus into a molecule of propylenediamine in place of two of its nitrogen
- atoms by addition and reduction reactions.
- 1 67. (Withdrawn): The method of Claim 66 wherein said step of incorporating comprises:
- preparing a first solution by dissolving propylenediamine into ethanol in a weight ratio of
- 3 about 1 to 50;

- admixing dimethylvinylphosphine sulfide into said solution in a weight ratio of about 1 to 22;
- 5 heating at reflux said solution for about 72 hours;
- 6 evaporating solvents under reduced pressure, leaving a residue.
- 1 68. (Withdrawn): The method of Claim 67 wherein said step of incorporating further
- 2 comprises:
- dissolving said residue in chloroform;
- washing said residue with NaOH; and
- 5 drying said residue over MgSO₄.
- 1 69. (original): The method of Claim 68 wherein said step of synthesizing further comprises:
- removing solvents in said residue under reduced pressure to yield an oil,
- 3 crystallizing said oil with ethyl acetate;
- preparing a suspension of LiAlH₄ in dry dioxane in a weight ratio of about 1 to 100;
- 5 admixing said oil into said suspension;
- 6 to yield a mixture;
- 7 refluxing said mixture for about 36 hours;
- 8 cooling said mixture; and
- adding a solution of dioxane in water and NaOH into said mixture.
- 1 70. (Withdrawn): The method of Claim 2 wherein said diseases consist of diabetes and
- abnormal low density lipoprotein (LDL) to high density lipoprotein (HDL) ratio and said
- 3 composition is selected from a group consisting of vanadyl 2,3,2-tetramine and chromium
- 4 2,3,2-tetramine; and
- said step of synthesizing further comprises reacting a metallic salt with 2,3,2-tetramine in an
- 6 ethanol solution.

- 71. (Withdrawn): The method of Claim 70 wherein said step of reacting comprises:
- 2 forming a first solution of 2,3,2 tetramine in ethanol in a weight ratio of about 1 to 20;
- forming a second solution of vanadyl acetylacetonate in ethanol in a weight ratio of about 1 to
- 4 275;
- 5 admixing said second solution into said first solution in a volume ratio of about 1 to 1; and
- 6 refluxing said solution for almost 30 minutes.
- 1 72. (Withdrawn): The method of Claim 70 wherein said step of reacting further comprises:
- 2 preparing a first solution of 2,3,2-tetramine in ethanol in a weight ratio of about 1 to 20;
- preparing a second solution of chromium (III) nitrate in ethanol in a weight ratio of about 1 to
- 4 80;
- admixing said second solution into said first solution in a volume ratio of about 1 to 1; and
- 6 refluxing said solution for about 30 minutes.
- 1 73. (Withdrawn): The method of Claim 55 wherein said step of converting comprises using
- amines to attach alkyl halide in a nucleophilic substitution of N atoms.
- 1 74. (previously presented): The method of Claim 3 wherein
- 2 said step of selecting comprises selecting a macrocyclic polyamine; and
- 3 said diseases comprise diabetes and diabetes-induced syndromes including congestive heart
- 4 failure, myocardial infarction, stroke, glaucoma, atherosclerosis, cardiomyopathy, ischemia,
- 5 optic neuropathy and peripheral neuropathy.
- 75. (previously presented): The method of claim 74 wherein said step of selecting comprises:

- ascertaining the heats of formation of a set of said compounds; and choosing said compound in consideration of its heat of formation compared to the heats of formation of other compounds in said set.
- 76. (previously presented): The method of claim 75 wherein: said step of ascertaining comprises: calculating the heats at the formation of said set of compounds from their respective constituent atoms.
- 77. (previously presented): The method of claim 76 wherein said step of choosing comprises determining the stabilities of said set of compounds as a function of their respective heats of formation;
- wherein said stabilities are determined in inverse proportion to said respective heats of formation; and
- whereby the relative stabilities of the set of compounds are deemed indicative of ability to yield the most stable complex when reacted with a group of metals.
- 1 78. (previously presented): The method of Claim 77 wherein;
- 2 said group of metals includes copper, cobalt, iron, zinc, cadmium, manganese and chromium.
- 79. (Withdrawn): The method of Claim 78 wherein said degenerative diseases comprise ischemic damage and pump failure post myocardial infarction characterized by iron-induced toxic redox effects and depletion of tissue zinc stores; and said compound is selected from a group consisting of zinc cyclam methylated, zinc cyclam adamantane, cyclam methylated and cyclam adamantane.

- 80. (Withdrawn): The method of claim 78 wherein said degenerative diseases comprise neurodegenerative disorders, stroke, glaucoma, atherosclerosis, cardiomyopathy, ischemia, optic neuropathy, peripheral neuropathy, presbycussis and cancer; and said composition is selected from derivatives of those compounds having the largest ring molecules.
- 81. (Withdrawn): The method of claim 80 wherein said compounds having the largest ring molecules includes 3,3,3 tetramine, cyclam adamantanes, cyclam 3,3,3 and compounds having alkyl substituted molecules.
- 82. (Withdrawn): The method of Claim 78 wherein said degenerative diseases comprise
 Parkinson's, Lou Gehrig's, Binswanger's, and Lewy Body diseases, Olivopontine Cerebellar
 Degeneration, stroke, glaucoma and optic neuropathy; and

said composition is selected from a group of compositions having alkyl side chains.

- 84. (Withdrawn): The method of claim 3 wherein said degenerative diseases comprise stroke, diabetic neuropathy, peripheral neuropathy, Alzheimer's disease, atherosclerosis, ischemia, diabetes, presbycussis, cardiomyopathy and congestive heart failure; and said composition is derived from compounds having terminal nitrogen added molecule substitution with elements selected from a group consisting of glutathione, uric acid, ascorbic acid, taurine, estrogen, dehydroepiandrosterone, probucol, vitamin E, hydroxytoluene, carvidilol, α lipoic acid, tocopherols, ubiquinone, phylloquinone, carotenes, menadione, glutamate, succinate, acetyl-l-carnitine, co-enzyme Q, lazeroids, polyphenolic flavonoids, homocysteine, menaquinone, idebenone, dantrolene and phosporous.
- 1 85. (Withdrawn): The method of Claim 84 wherein said degenerative disease comprises 2 stroke; and said composition consists of uric acid polyamine.

- 1 86. (Withdrawn): The method of Claim 84 wherein said degenerative disease comprises
- 2 diabetes; and said composition is derived from compounds selected from a group consisting of
- 3 phosphorous, taurine, CoEnzyme Q, α lipoic acid, tocopherol, succinate, glutamate and acetyl-
- 4 l-carnitine polyamines.
- 1 87. (Withdrawn): The method of Claim 84 wherein said degenerative disease comprises
- atherosclerosis; and said composition selected from a group consisting of tocopherol
- 3 polyamine and coenzyme Q polyamine.
- 1 88. (Withdrawn): The method of Claim 84 wherein said degenerative disease
- 2 comprises ischemia; and
- 3 said composition is selected from a group consisting of tocopherol polyamine and coenzyme Q
- 4 polyamine.
- 1 89. (Withdrawn): The method of Claim 84 wherein said diseases comprise myocardial
- degeneration and congestive heart failure; and said composition consists of coenzyme Q
- 3 polyamine.
- 90. (previously presented): The method of Claim 3 wherein said step of converting comprises
- adjusting the in vivo half life and pharmacokinetic properties of said composition by selective
- 3 terminal nitrogen substitutions.
- 91. (previously presented): The method of Claim 3 wherein said step of converting comprises
- adjusting the in vivo half life and pharmacokinetic properties of said composition by addition
- 3 of side chains on amino or methylene groups.

- 1 92. (previously presented): The method of Claim 3 wherein said step of selecting comprises:
- finding the octanol / water coefficients of partition of a series of said compounds; and
 - 3 picking said compound in consideration of its octanol / water coefficient compared to the
 - 4 octanol water coefficients of other compounds in said series.
 - 93. (previously presented): The method of Claim 92 wherein said step of picking comprises
 - determining the abilities of said series of compounds to pass through the intestinal, blood brain
 - and blood retinal barriers as a function of their respective octanol / water coefficients; wherein
 - said abilities are determined according to a distribution curve centered about 2 and having a
 - 5 useful range extending towards 0.5 and 4, the numbers being log values.
 - 1 94. (previously presented): The method of Claim 3 wherein said step of selecting comprises;
 - 2 measuring pKas of a list of said compounds; and
 - selecting said compound in consideration of its pKas compared to the pKa's of other
 - 4 compounds on the list.
 - 95. (previously presented): The method of Claim 94 wherein said step of selecting comprises;
 - selecting a composition with higher pKas in the treatment a disease characterized by lower
 - 3 tissue pH.
 - 1 96. (previously presented): The method of Claim 95 wherein said diseases include ischemia
 - 2 post myocardial infarction and diabetic ketoacidosis.

- 1 97. (previously presented): The method of Claim 3 wherein said step of selecting comprises
- determining the respective likely efficiency of said compounds in consideration of the disease
- 3 target to be treated and the route of administration.
- 1 98. (Withdrawn): The method of Claim 82 wherein said degenerative disease consists of
- 2 Alzheimer's disease and diabetes; and
- 3 said compound comprises acetyl-l-carnitine polyamine.
- 1 99. (Withdrawn): The method of Claim 84 wherein said degenerative disease consists of
- diabetes; and
- said compounds are selected from a group consisting of 2,3,2 piperidine, glutamate polyamine,
- succinate polyamine, chromium tetramine and vanadyl tetramine and phosphorous polyamine.
- 1 100. (Withdrawn): The method of Claim 3 wherein said degenerative diseases comprise
- 2 peripheral neuropathy and optic neuropathy; and
- said compounds comprise taurine polyamine and α lipoic acid polyamines.
- 1 101. (Withdrawn): The method of Claim 3 wherein said degenerative diseases comprise
- 2 glaucoma; and said compounds comprise adamantane 2,3,2 tetramine and adamantane cyclam.
- 1 102. (Withdrawn): The method of Claim 3 wherein said degenerative disease comprise
- presbycussis; and said compounds comprise α lipoic acid polyamine and acetyl-1-carnitine
- 3 polyamine.
- 1 103. (Withdrawn): The method of Claim 3 wherein said composition consists of:

- 2 1,4,8,11-tetraaza-1,4,8,11-tetramethylcyclotetradecane; and
- 3 said steps of synthesizing comprises:
- 4 refluxing for about 18 hours a solution of cyclam, formic acid, 37% formaldehyde and water
- in weight proportions of approximately 1, 5.3, 4.5 and 1 respectively;
- adding water to said solution in a weight ratio of approximately 0.5 to 1;
- cooling said solution to about 5°C;
- adjust the pH of said solution to above 12 with NaOH;
- 9 extracting the solution with CH₂Cl₂.
- 1 104. (Withdrawn): The method of Claim 2 wherein said composition consists of 1,4,8,11-
- tetraaza-1,4,8,11-tetra(2-piperidylethyl)cyclotetradecane; and said step of synthesizing further
- 3 comprises:
- preparing a first solution of cyclam and CH₂Cl₂ in a weight ratio of approximately 1 to 50;
- preparing a second solution of NaOH and water in a weight ratio of approximately 1 to 31;
- preparing a mixture of said first and second solution in a weight ratio of approximately 1 to 1;
- 7 preparing a third solution of 1-(2-chloroethyl)piperidine and CH₂Cl₂ in a weight ratio of
- 8 approximately 1 to 14;
- 9 adding said third solution dropwise into said mixture in a weight ratio of about 1 to 2;
- stirring said mixture over about 24 hours;
- evaporating solvents; and
- extracting residues with CH₂Cl₂.
- 1 105. (Withdrawn): The method of Claim 2 wherein said composition consists of 1,4,8,11-
- tetraaza-1,4,8,11 -tetrabicyclo[3.3.1]non-3-ylcyclotetradecane; and
- said step of synthesizing further comprises:
- forming a first solution of cyclam and ethanol in a weight ratio of approximately 1 to 100;

- forming a second solution of 1-bromoadamantane and ethanol in a weight ratio of 1 to 23;
- forming a mixture by adding said second solution dropwise into said first solution in a weight
- 7 ratio of about 1 to 1, over 30 minutes;
- 8 heating said mixture to reflux over about 20 hours;
- 9 evaporating said solution under reduced pressure; and
- 10 extracting residue from said solution with CH₂Cl₂.
- 1 106. (New): A method of treating degenerative diseases, said method comprising:
- 2 administering to a mammal an effective dose of non-superoxide dismutase mimic compound
- taken from a group consisting of those compounds having the formula
- 1 107. (Withdrawn): The method of Claim 78 wherein said degenerative diseases comprise
- 2 neurodegenerative diseases, ischemia post myocardial infarction and atherosclerosis; and
- said composition is selected from derivatives of compounds from a group consisting of
- 4 piperidine, piperazine and adamantane.

- 6 wherein A and B are hydrogen or alkyl, and m,n, and p are the same or different, and those
- 7 compounds having the formula

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